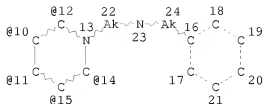
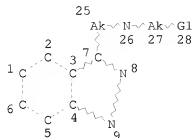


=> d 11
 L1 HAS NO ANSWERS
 L1 STR



VAR G1=10/11/12/14/15
 NODE ATTRIBUTES:
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 DEFAULT ECLEVEL IS LIMITED

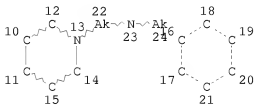
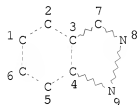
GRAPH ATTRIBUTES:
 RSPEC 16 13 6
 NUMBER OF NODES IS 28

STEREO ATTRIBUTES: NONE

=> d his 13

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 L3 0 S L1 FUL

=> d 16
 L6 HAS NO ANSWERS
 L6 STR



NODE ATTRIBUTES:
 DEFAULT MLEVEL IS ATOM
 DEFAULT ECLEVEL IS LIMITED

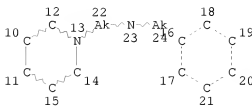
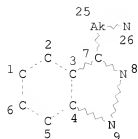
GRAPH ATTRIBUTES:
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 NUMBER OF NODES IS 24

STEREO ATTRIBUTES: NONE

=> d his 18

(FILE 'REGISTRY' ENTERED AT 16:23:41 ON 22 JUL 2008)
 L8 163 S L6 FUL

=> d 113
 L13 HAS NO ANSWERS
 L13 STR



NODE ATTRIBUTES:
 DEFAULT MLEVEL IS ATOM
 DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
 RSPEC 13 16 7
 NUMBER OF NODES IS 26

STEREO ATTRIBUTES: NONE

=> d his 114

(FILE 'REGISTRY' ENTERED AT 16:23:41 ON 22 JUL 2008)
 L14 2 SEARCH L13 SSS SUB=L8 FUL

=> fil caplus

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	ENTRY	SESSION
FULL ESTIMATED COST	489.92	490.34

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FILE COVERS 1907 - 22 Jul 2008 VOL 149 ISS 4
 FILE LAST UPDATED: 21 Jul 2008 (20080721/ED)

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<http://www.cas.org/legal/infopolicy.html>

=> s l14

L15 2 L14

=> d bib abs hitstr 1-2

L15 ANSWER 1 OF 2 CAPLUS COPYRIGHT 2008 ACS on SIN

AN 1999:246879 CAPLUS

DN 130:296684

TI Preparation of indazole- and 2-oxobenzamidazole-3-carboxamides as 5-HT₄ agonists and antagonists

IN Cohen, Marlene Lois; Schaus, John Mehnert; Thompson, Dennis Charles

PA Eli Lilly and Company, USA

SO Eur. Pat. Appl., 26 pp.

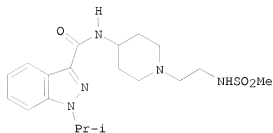
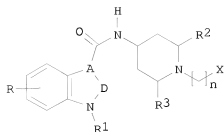
CODEN: EPXXDW

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	EP 908459	A1	19990414	EP 1998-308069	19981005
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, PT, IE, SI, LT, LV, FI, RO				
	US 6069152	A	20000530	US 1997-946495	19971007
	CA 2304826	A1	19990415	CA 1998-2304826	19980924
	WO 9917772	A1	19990415	WO 1998-US19992	19980924
	W: AL, AM, AT, AZ, BA, BB, BG, BR, BY, CA, CN, CU, CZ, EE, GD, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, RO, RU, SD, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
	JP 2001518504	T	20011016	JP 2000-514643	19980924
	US 6117882	A	20000912	US 1999-338707	19990623
PRAI	US 1997-946495	A	19971007		
	WO 1998-US19992	W	19980924		
OS	MARPAT 130:296684				
GI					



AB The title compds. [I; AD = C:N,NC:O; n = 1-5; R = H, halo, alkyl, etc.; R1 = H, alkyl, (un)substituted cycloalkyl; R2, R3 = H; R2R3 taken together form a bridge of 1-4 methylene units; X = OR4, NR4R5; R4 = H, alkyl, (un)substituted cycloalkyl, etc.; R5 = H; NR4R5 = pyrrolidino, piperazino, piperidino, etc.], antagonists and partial agonists for the serotonin receptor 5-HT4 which are useful for treatment of disorders caused by or affected by dysfunction of the 5-HT4 receptor such as anxiety, pain, depression, schizophrenia, memory disorders, dementia, irritable bowel syndrome, nausea, gastroesophageal reflux disease, dyspepsia, gastrointestinal motility disorders, constipation, atrial fibrillation, arrhythmias, tachycardia, urinary retention, urinary incontinence, or pain on urination, were prepared and formulated. E.g., methanesulfonylation of N-[1-(2-aminoethyl)piperidin-4-yl]-1-isopropylindazole-3-carboxamide (preparation given) afforded 60% II. Compds. I reduced the observed relaxations

of esophagus smooth muscle (of rats) at $\leq 10 \mu\text{M}$.

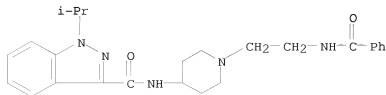
IT 207296-80-8P 207296-81-9P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of indazole- and 2-oxobenzamidazole-3-carboxamides as 5-HT4 agonists and antagonists)

RN 207296-80-8 CAPLUS

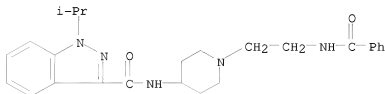
CN 1H-Indazole-3-carboxamide, N-[1-[2-(benzoylamino)ethyl]-4-piperidinyl]-1-(1-methylethyl)- (CA INDEX NAME)



RN 207296-81-9 CAPLUS
CN 1H-indazole-3-carboxamide, N-[1-[2-(benzoylamino)ethyl]-4-piperidinyl]-1-(1-methylethyl)-, ethanedioate (1:1) (CA INDEX NAME)

CM 1

CRN 207296-80-8
CMF C25 H31 N5 O2



CM 2

CRN 144-62-7
CMF C2 H2 O4



RE.CNT 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L15 ANSWER 2 OF 2 CAPLUS COPYRIGHT 2008 ACS on SIN

AN 1998:270001 CAPLUS

DN 128:316920

OREF 128:62633a

TI Synthesis and Structure-Activity Relationships of Potent and Orally Active 5-HT₄ Receptor Antagonists: Indazole and Benzimidazolone Derivatives

AU Schaus, John M.; Thompson, Dennis C.; Bloomquist, William E.; Susemichel, Alice D.; Calligaro, David O.; Cohen, Marlene L.

CS Lilly Research Laboratories, Eli Lilly and Company, Indianapolis, IN, 46285, USA

SO Journal of Medicinal Chemistry (1998), 41(11), 1943-1955

CODEN: JMCMAR; ISSN: 0022-2623

PB American Chemical Society

DT Journal

LA English

AB Indole-3-carboxamides, indazole-3-carboxamides, and benzimidazolone-3-carboxamides were synthesized and evaluated for antagonist affinity at the 5-HT₄ receptor in the rat esophagus. The endo-3-tropanamine derivs. in the indazole and benzimidazolone series possessed greater 5-HT₄ receptor affinity than the corresponding indole analogs. 5-HT₄ receptor antagonist affinity was further increased by alkylation at N-1 of the aromatic heterocycle. In 1-isopropylindazole-3-carboxamides, replacement of the bicyclic tropane ring system with the monocyclic piperidine ring system or an acyclic aminoalkylene chain led to potent 5-HT₄ receptor antagonists. In particular, those systems in which the basic amine was substituted with groups capable of forming H bonds showed increased 5-HT₄ receptor antagonist activity. While some of these compds. displayed high affinity

for other neurotransmitter receptors (in particular, 5-HT₃, α ₁, and 5-HT_{2A} receptors), as the conformational flexibility of the amine moiety increased, the selectivity for the 5-HT₄ receptor also increased. From this series of compds., the authors identified LY353433 (1-(1-methylethyl)-N-[2-[4-[(tricyclo[3.3.1.1^{3,7}]dec-1-ylcarbonyl)amino]-1-piperidinyl]ethyl]-1H-indazole-3-carboxamide) as a potent and selective 5-HT₄ receptor antagonist with clin. suitable pharmacodynamics.

IT 207296-81-9P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(synthesis and structure-activity relationships of potent and orally active indazole and benzimidazolone 5-HT₄ receptor antagonists)

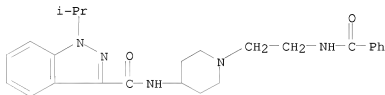
RN 207296-81-9 CAPLUS

CN 1H-Indazole-3-carboxamide, N-[1-[2-(benzoylamino)ethyl]-4-piperidinyl]-1-(1-methylethyl)-, ethanedioate (1:1) (CA INDEX NAME)

CM 1

CRN 207296-80-8

CMF C25 H31 N5 O2



CM 2

CRN 144-62-7

CMF C2 H2 O4



RE.CNT 37 THERE ARE 37 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT